

What is claimed is:

- 1        1. A method for reconstructing an image of a scattering medium, comprising:
  - 2              directing energy into the scattering medium at a source location on the
  - 3              scattering medium;
  - 4              measuring the energy emerging from the scattering medium at a detector
  - 5              location on the scattering medium;
  - 6              selecting an initial guess of internal properties of the scattering medium;
  - 7              predicting the energy emerging from the scattering medium using an
  - 8              equation of radiative transfer, wherein the prediction is a function of the initial guess;
  - 9              generating an objective function based on a comparison of the prediction
  - 10          with the measurement;
  - 11          generating a gradient of the objective function by a method of adjoint
  - 12          differentiation;
  - 13          modifying the initial guess of the properties of the scattering medium
  - 14          based on the gradient of the objective function; and
  - 15          generating an image representation of the internal properties of the
  - 16          scattering medium.

- 1        2. The method according to claim 1, further comprising repeating the
- 2          predicting of the energy emerging from the scattering medium based on the modified
- 3          initial guess, generating the objective function and modifying the initial guess, until at
- 4          least one of a predetermined number of repetitions has occurred and the objective
- 5          function reaches a predetermined threshold.

1       3.     The method according to claim 1, wherein the prediction depends on the  
2     boundary conditions.

1       4.     The method according to claim 3, wherein the boundary conditions  
2     account for a refractive mismatch at an interface between the medium and at least one of  
3     the detectors and source.

1       5.     The method according to claim 1, wherein the prediction comprises an  
2     iterative process producing intermediate results.

1       6.     The method according to claim 5, wherein the intermediate results of the  
2     prediction are stored.

1       7.     The method according to claim 6, wherein generating the gradient of the  
2     objective function by adjoint differences uses the intermediate results of the prediction.

1       8.     The method according to claim 7, wherein generating the gradient  
2     comprises stepping backward through the intermediate results of the prediction.

1       9.     The method according to claim 1, wherein the equation of radiative  
2     transfer is time independent.

1       10.    The method according to claim 9, wherein the time independent equation  
2     of radiative transfer is:

3                    $\omega \nabla \Psi(\mathbf{r}, \omega) + (\mu_a + \mu_s) \Psi(\mathbf{r}, \omega) = S(\mathbf{r}, \omega) + \mu_s \int_0^{2\pi} p(\omega, \omega') \Psi(\mathbf{r}, \omega') d\omega'$

4                   where  $\Psi(\mathbf{r}, \omega)$  is the radiance at the spatial position  $\mathbf{r}$  directed into a unit  
5                   solid angle  $\omega$ ,  $S(\mathbf{r}, \omega)$  is the energy directed into the medium at spatial position  $\mathbf{r}$  into a  
6                   unit solid angle  $\omega$ ,  $\mu_s$  is the scattering coefficient,  $\mu_a$  is the absorption coefficient and  
7                    $p(\omega, \omega')$  is the scattering phase function.

1                   11.       The method according to claim 10, wherein the scattering phase function  
2                   is:

3                   
$$p(\cos\theta) = \frac{1 - g^2}{2(1 + g^2 - 2g\cos\theta)^{3/2}}$$

4                   where  $\theta$  is the angle between the two unit solid angles  $\omega$  and  $\omega'$ , and  $g$  is  
5                   the anisotropy factor.

1                   12.       The method according to claim 1, wherein the equation of radiative  
2                   transfer is time dependent.

1                   13.       The method according to claim 12, wherein the time dependent equation  
2                   of radiative transfer is:

3                   
$$\frac{1}{c} \frac{\partial \Psi(\mathbf{r}, \omega, t)}{\partial t} = S(\mathbf{r}, \omega, t) - \omega \cdot \nabla \Psi(\mathbf{r}, \omega, t) - (\mu_a + \mu_s) \Psi(\mathbf{r}, \omega, t) + \mu_s \int_0^{2\pi} p(\omega, \omega') \Psi(\mathbf{r}, \omega', t) d\omega'$$

4                   where  $\Psi(\mathbf{r}, \omega, t)$  is the radiance at the spatial position  $\mathbf{r}$  directed into a unit  
5                   solid angle  $\omega$ ,  $S(\mathbf{r}, \omega, t)$  is the energy directed into the medium at spatial position  $\mathbf{r}$  into a

6 unit solid angle  $\omega$ ,  $\mu_s$  is the scattering coefficient,  $\mu_a$  is the absorption coefficient and  
7  $p(\omega, \omega')$  is the scattering phase function.

1 14. The method according to claim 13, wherein the scattering phase function  
2 is:

3 
$$p(\cos\theta) = \frac{1 - g^2}{2(1 + g^2 - 2g\cos\theta)^{3/2}}$$

4 where  $\theta$  is the angle between the two unit solid angles  $\omega$  and  $\omega'$ , and  $g$  is  
5 the anisotropy factor.

1 15. The method according to claim 1, wherein the properties include at least  
2 one of a scattering coefficient, an absorption coefficient, an anisotropy factor, and a  
3 scattering phase function.

1 16. The method according to claim 1, wherein the objective function is a  
2 normalized comparison of the predicted energy and the measured energy

1 17. The method according to claim 1, wherein the objective function is based  
2 on the normalized sum of the differences between the predicted energy and the measured  
3 energy for each source detector pair, wherein a source detector pair is formed between  
4 each source location and each detector location.

1 18. The method according to claim 1, wherein the objective function is:

2                    $\varphi = \frac{1}{2} \sum_i^m (P_i - M_i)^2$

3                   where  $M_i$  represents the actual measurements and the  $P_i$  represents the  
4                   predicted measurements for each source defector pair  $i$ ,  $m$  is the number of source  
5                   detector pairs, where a source detector pairs is formed between each source location and  
6                   each detector location.

1                   19.       The method according to claim 1, further comprising minimizing the  
2                   objective function.

1                   20.       The method according to claim 19, wherein minimizing the objective  
2                   function includes a one dimensional line search.

1                   21.       The method according to claim 20, wherein the one dimensional line  
2                   search is performed along a direction of the gradient of the objective function.

1                   22.       The method according to claim 20, wherein the one dimensional line  
2                   search is performed along a gradient-dependent direction.

1                   23.       The method according to claim 1, wherein the energy comprises near  
2                   infra-red energy.

1        24. The method according to claim 1, wherein the scattering medium contains  
2 regions wherein the scattering coefficients are not substantially greater than the  
3 absorption coefficients.

1        25. The method according to claim 1, wherein the scattering medium contains  
2 a low scattering region embedded in a high scattering region.

1        26. The method according to claim 1, wherein the predicted energy is  
2 determined using finite element methods.

1        27. The method according to claim 1, wherein the predicted energy is  
2 determined using finite difference methods.

1        28. A method for imaging the spatial optical properties of tissue, comprising:  
2            (a) directing energy into the scattering medium at a source location on  
3 the tissue;  
4            (b) measuring the energy emerging from the scattering medium at a  
5 detector location on the tissue;  
6            (c) selecting and initial guess of the spatial optical properties of the  
7 tissue;  
8            (d) predicting the energy emerging from the tissue using an equation  
9 of radiative transfer in an iterative process, wherein the prediction is a function of the

10 initial guess and a refraction index mismatch at a boundary of the tissue, and the iterative  
11 process generates a plurality of intermediate predictions;

12 (e) generating an objective function based on a normalized

13 comparison of the prediction with the measured energy emerging from the scattering  
14 medium;

15 (f) generating a gradient of the objective function by adjoint  
16 differentiation;

17 (g) modifying the initial guess of the spatial properties of the tissue  
18 based on the gradient of the objective function;

19 (h) repeating steps (d) through (g) until at least one of a threshold of  
20 modifications to the initial guess is reached and the objective function reaches a  
21 threshold; and

22 (j) generating an image representation of the spatial optical properties  
23 of the tissue.

1 29. A system for reconstructing an image of a scattering medium, comprising:

2 a source for directing energy into the scattering medium at source location on the  
3 scattering medium;

4 a detector for measuring the energy emerging from the scattering medium at a  
5 detector location on the scattering medium;

6 an initial guess of internal properties of the scattering medium;

7 means for predicting the energy emerging from the scattering medium using an  
8 equation of radiative transfer, wherein the prediction is a function of the initial guess;

9           means for generating an objective function based on a comparison of the  
10          prediction with the measurement;  
11           means for generating a gradient of the objective function by a method of adjoint  
12          differentiation;  
13           means for modifying the initial guess of the properties of the scattering medium  
14          based on the gradient of the objective function; and  
15           means for generating an image representation of the internal properties of the  
16          scattering medium.

1           30.       The system according to claim 1, further comprising means for repeating  
2          the predicting of the energy emerging from the scattering medium based on the modified  
3          initial guess, generating the objective function and modifying the initial guess, until at  
4          least one of a predetermined number of repetitions has occurred and the objective  
5          function reaches a predetermined threshold.

1           31.       The system according to claim 1, wherein the prediction depends on the  
2          boundary conditions.

1           32.       The system according to claim 31, wherein the boundary conditions  
2          account for a refractive mismatch at an interface between the medium and at least one of  
3          the detectors and source.

1        33. The system according to claim 1, wherein the prediction comprises an  
2 iterative process producing intermediate results.

1        34. The system according to claim 33, wherein the intermediate results of the  
2 prediction are stored.

1        35. The system according to claim 34, wherein generating the gradient of the  
2 objective function by adjoint differences uses the intermediate results of the prediction.

1        36. The system according to claim 35, wherein generating the gradient  
2 comprises stepping backward through the intermediate results of the prediction.

1        37. The system according to claim 1, wherein the equation of radiative  
2 transfer is time independent.

1        38. The system according to claim 37, wherein the time independent equation  
2 of radiative transfer is:

3         $\omega \nabla \Psi(\mathbf{r}, \omega) + (\mu_a + \mu_s) \Psi(\mathbf{r}, \omega) = S(\mathbf{r}, \omega) + \mu_s \int_0^{2\pi} p(\omega, \omega') \Psi(\mathbf{r}, \omega') d\omega'$

4        where  $\Psi(\mathbf{r}, \omega)$  is the radiance at the spatial position  $\mathbf{r}$  directed into a unit solid  
5 angle  $\omega$ ,  $S(\mathbf{r}, \omega)$  is the energy directed into the medium at spatial position  $\mathbf{r}$  into a unit  
6 solid angle  $\omega$ ,  $\mu_s$  is the scattering coefficient,  $\mu_a$  is the absorption coefficient and  $p(\omega, \omega')$   
7 is the scattering phase function.

1        39. The system according to claim 38, wherein the scattering phase function  
2        is:

3                      
$$p(\cos\theta) = \frac{1-g^2}{2(1+g^2 - 2g\cos\theta)^{3/2}}$$

4        where  $\theta$  is the angle between the two unit solid angles  $\omega$  and  $\omega'$ , and  $g$  is the  
5        anisotropy factor.

1        40. The system according to claim 1, wherein the equation of radiative  
2        transfer is time dependent.

1        41. The system according to claim 40, wherein the time dependent equation of  
2        radiative transfer is:

3                      
$$\frac{1}{c} \frac{\partial \Psi(r,\omega,t)}{\partial t} = S(r,\omega,t) - \omega \cdot \nabla \Psi(r,\omega,t) - (\mu_a + \mu_s) \Psi(r,\omega,t) + \mu_s \int_0^{2\pi} p(\omega,\omega') \Psi(r,\omega',t) d\omega'$$
  
4        where  $\Psi(r,\omega,t)$  is the radiance at the spatial position  $r$  directed into a unit solid  
5        angle  $\omega$ ,  $S(r,\omega,t)$  is the energy directed into the medium at spatial position  $r$  into a unit  
6        solid angle  $\omega$ ,  $\mu_s$  is the scattering coefficient,  $\mu_a$  is the absorption coefficient and  $p(\omega,\omega')$   
7        is the scattering phase function.

1        42. The system according to claim 41, wherein the scattering phase function  
2        is:

3                      
$$p(\cos\theta) = \frac{1-g^2}{2(1+g^2 - 2g\cos\theta)^{3/2}}$$

4        where  $\theta$  is the angle between the two unit solid angles  $\omega$  and  $\omega'$ , and  $g$  is the  
5        anisotropy factor.

1            43.      The system according to claim 1, wherein the properties include at least  
2        one of a scattering coefficient, an absorption coefficient, an anisotropy factor, and a  
3        scattering phase function.

1            44.      The system according to claim 1, wherein the objective function is a  
2        normalized comparison of the predicted energy and the measured energy.

1            45.      The system according to claim 1, wherein the objective function is based  
2        on the normalized sum of the differences between the predicted energy and the measured  
3        energy for each source detector pair, wherein a source detector pair is formed between  
4        each source location and each detector location.

1            46.      The system according to claim 1, wherein the objective function is:

2

$$\varphi = \frac{1}{2} \sum_i^m (P_i - M_i)^2$$

3        where  $M_i$  represents the actual measurements and  $P_i$  represents the predicted  
4        measurements for each source detector pair,  $m$  is the number of source detector pairs,  
5        where a source detector pairs is formed between each source location and each detector  
6        location.

1        47. The system according to claim 1, further comprising minimizing the  
2 objective function.

1        48. The system according to claim 47, wherein minimizing the objective  
2 function includes a one dimensional line search.

1        49. The system according to claim 48, wherein the one dimensional line  
2 search is performed along a direction of the gradient of the objective function.

1        50. The system according to claim 49, wherein the one dimensional line  
2 search is performed along a gradient-dependent direction.

1        51. The system according to claim 50, wherein the energy comprises near  
2 infra-red energy.

1        52. The system according to claim 1, wherein the scattering medium contains  
2 regions wherein the scattering coefficients are not substantially greater than the  
3 absorption coefficients.

1        53. The system according to claim 1, wherein the scattering medium contains  
2 a low scattering region embedded in a high scattering region.

1        54. The system according to claim 1, wherein the predicted energy is  
2 determined using finite element methods.

1        55. The system according to claim 1, wherein the predicted energy is  
2 determined using finite difference methods.

1        56. A system for imaging the spatial distribution of optical properties of  
2 tissue, comprising:

3            (a) a source for directing energy into the scattering medium at a source  
4 location on the tissue;

5            (b) a detector for measuring the energy emerging from the scattering medium  
6 at a detector location on the tissue;

7            (c) an initial guess of spatial optical properties of the tissue;

8            (d) means for predicting the energy emerging from the tissue using an  
9 equation of radiative transfer in an iterative process, wherein the prediction is a function  
10 of the initial guess and a refraction index mismatch at a boundary of the tissue, and the  
11 iterative process generates a plurality of intermediate predictions;

12            (e) means for generating an objective function based on a normalized  
13 comparison of the prediction with the measured energy emerging from the scattering  
14 medium;

15            (f) means for generating a gradient of the objective function by adjoint  
16 differentiation;

17           (g)     means for modifying the initial guess of the spatial properties of the tissue  
18     based on the gradient of the objective function;  
19           (h)     means for repeating steps (d) through (g) until at least one of a threshold  
20     of modifications to the initial guess is reached and the objective function reaches a  
21     threshold; and  
22           (j)     means for generating an image representation of the spatial optical  
23     properties of the tissue.

1               57.    Computer executable software code stored on a computer readable  
2     medium, the code for reconstructing an image of a scattering medium, comprising:  
3                      code to direct energy into the scattering medium at a source location on  
4     the scattering medium;  
5                      code to measure the energy emerging from the scattering medium at a  
6     detector location on the scattering medium;  
7                      code to receive an initial guess of internal properties of the scattering  
8     medium;  
9                      code to predict the energy emerging from the scattering medium using an  
10    equation of radiative transfer, wherein the prediction is a function of the initial guess;  
11                      code to generate an objective function based on a comparison of the  
12    prediction with the measurement;  
13                      code to generate a gradient of the objective function by a method of  
14    adjoint differentiation;

15 code to modify the initial guess of the properties of the scattering medium  
16 based on the gradient of the objective function; and  
17 code to generate an image representation of the internal properties of the  
18 scattering medium.

1 58. Computer executable software code stored on a computer readable  
2 medium, the code for imaging the spatial distribution of optical properties of tissue,  
3 comprising:

- 4 (a) code to direct energy into the scattering medium at a source location on the  
5 tissue;  
6 (b) code to measure the energy emerging from the scattering medium at a detector  
7 location on the tissue;  
8 (c) code to receive an initial guess of spatial optical properties of the tissue;  
9 (d) code to predict the energy emerging from the tissue using an equation of  
10 radiative transfer in an iterative process, wherein the prediction is a function of the initial  
11 guess and a refraction index mismatch at a boundary of the tissue, and the iterative  
12 process generates a plurality of intermediate predictions;  
13 (e) code to generate an objective function based on a normalized comparison of  
14 the prediction with the measured energy emerging from the scattering medium;  
15 (f) code to generate a gradient of the objective function by adjoint differentiation;  
16 (g) code to modify the initial guess of the spatial properties of the tissue based on  
17 the gradient of the objective function;

18           (h) code to repeat steps (d) through (g) until at least one of a threshold of  
19        modifications to the initial guess is reached and the objective function reaches a  
20        threshold; and  
21           (j) code to generate an image representation of the spatial optical properties of the  
22        tissue.

1           59.     A computer readable medium having computer executable software code  
2        stored thereon, the code for reconstructing an image of a scattering medium, comprising:  
3                  code to direct energy into the scattering medium at a source location on  
4        the scattering medium;  
5                  code to measure the energy emerging from the scattering medium at a  
6        detector location on the scattering medium;  
7                  code to receive an initial guess of internal properties of the scattering  
8        medium;  
9                  code to predict the energy emerging from the scattering medium using an  
10      equation of radiative transfer, wherein the prediction is a function of the initial guess;  
11                  code to generate an objective function based on a comparison of the  
12      prediction with the measurement;  
13                  code to generate a gradient of the objective function by a method of  
14      adjoint differentiation;  
15                  code to modify the initial guess of the properties of the scattering medium  
16      based on the gradient of the objective function; and

17 code to generate an image representation of the internal properties of the  
18 scattering medium.

1 60. A computer readable medium having computer executable software code  
2 stored thereon, the code for imaging the spatial distribution of optical properties of tissue,  
3 comprising:

4 (a) code to direct energy into the scattering medium at a source location on the  
5 tissue;

6 (b) code to measure the energy emerging from the scattering medium at a detector  
7 location on the tissue;

8 (c) code to receive an initial guess of spatial optical properties of the tissue;

9 (d) code to predict the energy emerging from the tissue using an equation of  
10 radiative transfer in an iterative process, wherein the prediction is a function of the initial  
11 guess and a refraction index mismatch at a boundary of the tissue, and the iterative  
12 process generates a plurality of intermediate predictions;

13 (e) code to generate an objective function based on a normalized comparison of  
14 the prediction with the measured energy emerging from the scattering medium;

15 (f) code to generate a gradient of the objective function by adjoint differentiation;

16 (g) code to modify the initial guess of the spatial properties of the tissue based on  
17 the gradient of the objective function;

18 (h) code to repeat steps (d) through (g) until at least one of a threshold of  
19 modifications to the initial guess is reached and the objective function reaches a  
20 threshold; and

21           (j) code to generate an image representation of the spatial optical properties of the  
22           tissue.

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